

Access DB# 84663**SEARCH REQUEST FORM**

Scientific and Technical Information Center

**JAN**

Requester's Full Name: FONDA Examiner #: 71970 Date: 1-17-03  
 Art Unit: 1623 Phone Number 30 8-1620 Serial Number: 09/760879  
 Mail Box and Bldg/Room Location: CHI 8B19 CHI 8A05 Results Format Preferred (circle): PAPER DISK E-MAIL

If more than one search is submitted, please prioritize searches in order of need.

\*\*\*\*\*  
 Please provide a detailed statement of the search topic, and describe as specifically as possible the subject matter to be searched. Include the elected species or structures, keywords, synonyms, acronyms, and registry numbers, and combine with the concept or utility of the invention. Define any terms that may have a special meaning. Give examples or relevant citations, authors, etc, if known. Please attach a copy of the cover sheet, pertinent claims, and abstract.

Title of Invention: \_\_\_\_\_

Inventors (please provide full names): \_\_\_\_\_

Earliest Priority Filing Date: 5-19-99

\*For Sequence Searches Only\* Please include all pertinent information (parent, child, divisional, or issued patent numbers) along with the appropriate serial number.

Please search compositions comprising hyaluronic acid bonded to an agent for treating joint disease, as in claims 1-10, 12-14, and 18-21. Please also search preparative method of claim 11 and therapeutic methods of claims 17 and 22.

Thanks.

K.

Jan Delaval  
 Reference Librarian  
 Biotechnology & Chemical Library  
 CM1 1E07 - 703-308-4498  
 jan.delaval@uspto.gov

**STAFF USE ONLY**

Searcher: Jan  
 Searcher Phone #: 4458  
 Searcher Location: \_\_\_\_\_  
 Date Searcher Picked Up: 1/21/03  
 Date Completed: 1/24/03  
 Searcher Prep & Review Time: \_\_\_\_\_  
 Clerical Prep Time: 15  
 Online Time: 4120

**Type of Search**

NA Sequence (#) \_\_\_\_\_  
 AA Sequence (#) \_\_\_\_\_  
 Structure (#) ✓  
 Bibliographic ✓  
 Litigation \_\_\_\_\_  
 Fulltext \_\_\_\_\_  
 Patent Family \_\_\_\_\_  
 Other \_\_\_\_\_

**Vendors and cost where applicable**

STN ✓  
 Dialog \_\_\_\_\_  
 Questel/Orbit \_\_\_\_\_  
 Dr.Link \_\_\_\_\_  
 Lexis/Nexis \_\_\_\_\_  
 Sequence Systems ✓  
 WWW/Internet \_\_\_\_\_  
 Other (specify) \_\_\_\_\_

(FILE 'HOME' ENTERED AT 15:52:25 ON 21 JAN 2003)  
SET COST OFF

FILE 'REGISTRY' ENTERED AT 15:52:46 ON 21 JAN 2003

L1	2 S	HYALURONIC ACID/CN OR 9067-32-7
L2	753 S	?HYALURON?/CNS NOT' L1
L3	435 S	L2 NOT SQL/FA
L4	318 S	L2 NOT L3
	E	CYCLOOXYGENASE/CN
L5	1 S	E8
L6	2 S	E3, E7
	E	MATRIX METALLOPROTEASE/CN
L7	15 S	E3, E5-E13, E15-E17, E23, E24
L8	5 S	E25, E36, E43, E45, E46
L9	4 S	E50, E51, E55, E58
L10	1 S	E61
L11	5 S	E72, E75, E79-E81
L12	4 S	E85, E89-E91
L13	1365 S	(?METALLOPROTEINASE? OR ?METALLOPROTEASE?)/CNS
L14	STR	
L15	31 S	L14 CSS
L16	2264 S	L14 FUL
	SAV TEMP	L16 FONDA700/A
L17	629 S	L14 CSS FUL SUB=L16
	SAV	L17 FONDA700A/A

Jan Delaval  
Reference Librarian  
Biotechnology & Chemical Library  
CM1 1E07 - 703-308-4498  
jan.delaval@uspto.gov

FILE 'HCAPLUS' ENTERED AT 16:16:23 ON 21 JAN 2003

L18	10031 S	L1
L19	3440 S	L3
L20	151 S	L4
L21	14614 S	HYALURONIC ACID OR HYALURONATE OR HYALURONAN
L22	20161 S	?HYALURON?
L23	20696 S	L18-L22
L24	1922 S	L5
L25	9113 S	L6
L26	13384 S	(COX OR CYCLOOXYGENASE OR CYCLO OXYGENASE) (L)2 OR COX2
L27	13 S	PROSTAGLANDIN(L) (ENDOPEROXIDASE OR ENDO PEROXIDASE) (L) (SYNTHA
L28	41 S	L23 AND L24-L27
L29	26594 S	L7-L13
L30	476 S	L23 AND L29
L31	309 S	L17
L32	4 S	L23 AND L31

FILE 'REGISTRY' ENTERED AT 16:21:16 ON 21 JAN 2003

L33	1635 S	L16 NOT L17
-----	--------	-------------

FILE 'HCAPLUS' ENTERED AT 16:21:22 ON 21 JAN 2003

L34	3 S	L33 AND L23
L35	45 S	L28, L32, L34
	E	ANTIRHEUMAT/CT
	E	E5+ALL
L36	4437 S	E5, E4+NT
L37	48 S	L23 AND L36
L38	91 S	L35, L37
L39	77 S	L23 AND (ANTIRHEUMAT? OR ANTI RHEUMAT?)
L40	136 S	L38, L39
L41	6 S	L40 AND ?CONJUGAT?
	E	TAMURA T/AU
L42	596 S	E3-E5
	E	TAMURA TATSUYA/AU

=> d his

(FILE 'REGISTRY' ENTERED AT 16:36:53 ON 27 JAN 2003)  
DEL HIS  
L1 2 S HYALURONIC ACID/CN OR 9067-32-7  
L2 773 S ?HYALURON?/CNS  
ACT FONDA700A/A  
-----  
L3 STR  
L4 ( 2264)SEA FILE=REGISTRY SSS FUL L3  
L5 629 SEA FILE=REGISTRY SUB=L4 CSS FUL L3  
-----

FILE 'HCAPLUS' ENTERED AT 16:39:55 ON 27 JAN 2003  
L6 10046 S L1  
L7 12812 S L2  
L8 14642 S HYALURONIC ACID OR HYALURONATE OR HYALURONAN  
L9 20190 S ?HYALURON?  
L10 20726 S L6-L9  
L11 310 S L5  
L12 4 S L11 AND L10

FILE 'HCAPLUS' ENTERED AT 16:40:56 ON 27 JAN 2003

=> fil hcaplus

FILE 'HCAPLUS' ENTERED AT 16:41:26 ON 27 JAN 2003  
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.  
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.  
COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 27 Jan 2003 VOL 138 ISS 5  
FILE LAST UPDATED: 26 Jan 2003 (20030126/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d l12 all hitstr tot

L12 ANSWER 1 OF 4 HCAPLUS COPYRIGHT 2003 ACS  
AN 2002:716020 HCAPLUS  
DN 137:293053  
TI Medical devices and compositions for treating vulnerable plaque  
IN Brown, David L.  
PA Volcano Therapeutics, Inc., USA  
SO PCT Int. Appl., 28 pp.  
CODEN: PIXXD2  
DT Patent  
LA English  
IC ICM A61K  
CC 63-7 (Pharmaceuticals)  
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2002072014	A2	20020919	WO 2002-US7244	20020308
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
	US 2003004141	A1	20030102	US 2002-96131	20020308
PRAI	US 2001-274331P	P	20010308		
AB	Medical devices, compns. and methods for treating or preventing atherosclerotic plaque rupture are disclosed. Specifically, medical devices that deliver to a treatment site metalloproteinase inhibitors (MMPI) are disclosed. The medical devices include catheters, guide wires, vascular stents, micro-particles, electronic leads, probes, sensors, drug depots, transdermal patches, and vascular patches. Representative MMPIs included zinc chelators, urea derivs., caprolactone-based inhibitors, phosphonamides, piperazines, sulfonamides, tertiary amines, carbamate derivs., mercapto alcs., mercapto ketones, antimicrobial tetracyclines, non-antimicrobial tetracyclines, and derivs. and combinations thereof. In one embodiment a self-expanding vascular stent is coated with at least one MMPI and deployed at a site within an artery where vulnerable plaque has been identified.				
ST	medical device plaque; polymer coating medical device plaque; drug				
	delivery medical device plaque				
IT	Polyesters, biological studies				
	RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (caprolactone-based; medical devices and compns. for treating vulnerable plaque)				
IT	Medical goods				
	(catheters; medical devices and compns. for treating vulnerable plaque)				
IT	Drug delivery systems				
	(controlled-release; medical devices and compns. for treating vulnerable plaque)				
IT	Polyesters, biological studies				
	RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (dilactone-based; medical devices and compns. for treating vulnerable plaque)				
IT	Polyesters, biological studies				
	RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (hydroxycarboxylic acid-based; medical devices and compns. for treating vulnerable plaque)				
IT	Polyesters, biological studies				
	RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (lactic acid-based; medical devices and compns. for treating vulnerable plaque)				
IT	Cellophane				
	Coating materials				
	Electric contacts				
	Human				
	Medical goods				
	Sensors				
	(medical devices and compns. for treating vulnerable plaque)				
IT	Acrylic polymers, biological studies				

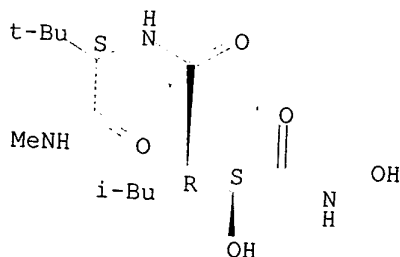
Alkyd resins  
 Collagens, biological studies  
 Epoxy resins, biological studies  
 Fibrinogens  
 Fibrins  
 Fluoropolymers, biological studies  
 Polyamides, biological studies  
 Polyanhydrides  
 Polycarbonates, biological studies  
 Polyesters, biological studies  
 Polyethers, biological studies  
 Polyimides, biological studies  
 Polyolefins  
 Polyoxymethylenes, biological studies  
 Polyphosphazenes  
 Polysiloxanes, biological studies  
 Polyurethanes, biological studies  
 Rayon, biological studies  
 RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (medical devices and compns. for treating vulnerable plaque)  
 IT Sulfonamides  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (medical devices and compns. for treating vulnerable plaque)  
 IT Alcohols, biological studies  
 Ketones, biological studies  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (mercapto; medical devices and compns. for treating vulnerable plaque)  
 IT Polyethers, biological studies  
 RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (ortho ester group-contg.; medical devices and compns. for treating vulnerable plaque)  
 IT Tooth  
 (plaque; medical devices and compns. for treating vulnerable plaque)  
 IT Polyethers, biological studies  
 RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (polyester-; medical devices and compns. for treating vulnerable plaque)  
 IT Polyesters, biological studies  
 RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (polyether-; medical devices and compns. for treating vulnerable plaque)  
 IT Vinyl compounds, biological studies  
 RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (polymers; medical devices and compns. for treating vulnerable plaque)  
 IT Medical goods  
 (stents; medical devices and compns. for treating vulnerable plaque)  
 IT Amines, biological studies  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (tertiary; medical devices and compns. for treating vulnerable plaque)  
 IT Drug delivery systems  
 (transdermal; medical devices and compns. for treating vulnerable plaque)  
 IT Medical goods  
 (wires; medical devices and compns. for treating vulnerable plaque)  
 IT 9001-12-1, Metalloproteinase-1  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (inhibitors; medical devices and compns. for treating vulnerable plaque)

- IT 9002-85-1, Polyvinylidene chloride 9002-86-2, PVC 9003-09-2, Polyvinyl methyl ether 9003-20-7, Poly(vinyl acetate) 9003-53-6, Polystyrene 9003-54-7, Acrylonitrile-styrene copolymer 9003-56-9, Acrylonitrile-butadiene-styrene copolymer 9003-63-8, Poly(butyl methacrylate) 9004-32-4, Carboxymethyl cellulose 9004-34-6, Cellulose, biological studies 9004-35-7, Cellulose acetate 9004-36-8, Cellulose acetate butyrate 9004-48-2, Cellulose propionate 9004-61-9, **Hyaluronic acid** 9004-70-0, Cellulose nitrate 9005-25-8, Starch, biological studies 9015-12-7, Cellulose butyrate 24937-78-8, EVA 24937-79-9, Polyvinylidene fluoride 24980-41-4, Polycaprolactone 25014-41-9, Polyacrylonitrile 25038-54-4, Polycaprolactam, biological studies 25101-13-7, Ethylene-methyl methacrylate copolymer 25248-42-4, Polycaprolactone 25249-16-5, Poly(2-hydroxyethyl methacrylate) 26009-03-0, PolyGlycolic acid 26023-30-3, Poly[oxy(1-methyl-2-oxo-1,2-ethanediyl)] 26100-51-6, Poly(DL-lactic acid) 26124-68-5, PolyGlycolic acid 26161-42-2 26780-50-7, Glycolide-lactide copolymer 26811-96-1, Poly(L-lactic acid) 29223-92-5 31621-87-1, Polydioxanone 31852-84-3, Poly(trimethylene carbonate) 32131-17-2, Nylon 66, biological studies 50862-75-4, Poly(oxy carbonyloxy-1,3-propanediyl) 113883-69-5, Glycolic acid-trimethylene carbonate copolymer 128171-16-4, Hydroxybutyric acid-hydroxyvaleric acid copolymer
- RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
- (medical devices and compns. for treating vulnerable plaque)
- IT 57-13-6D, Urea, derivs. 60-54-8, Tetracycline 110-85-0D, Piperazine, derivs. 463-77-4D, Carbamic acid, derivs. 502-44-3D, Caprolactone, derivs. 564-25-0, Doxycycline 7440-66-6D, Zinc, chelates 10118-90-8, Minocycline 88828-25-5, CMT 8 130370-60-4, Batimastat 154039-60-8, Marimastat
- RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
- (medical devices and compns. for treating vulnerable plaque)
- IT **9004-61-9, Hyaluronic acid**
- RL: DEV (Device component use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
- (medical devices and compns. for treating vulnerable plaque)
- RN 9004-61-9 HCAPLUS
- CN Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

- IT **154039-60-8, Marimastat**
- RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
- (medical devices and compns. for treating vulnerable plaque)
- RN 154039-60-8 HCAPLUS
- CN Butanediamide, N4-[(1S)-2,2-dimethyl-1-[(methylamino)carbonyl]propyl]-N1,2-dihydroxy-3-(2-methylpropyl)-, (2S,3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



X

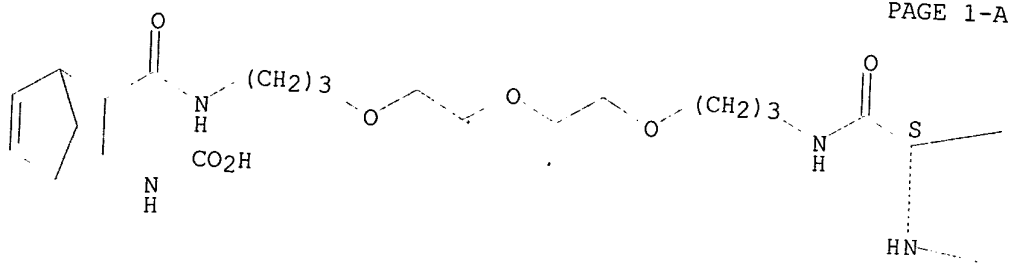
[illegible]

AB Disclosed is a compd. having MMP inhibitory activity which is a compd. of a hydroxamic acid deriv. I and **hyaluronic acid**, wherein R1 = H, OH, C1-8 alkyl, etc.; R2 = C1-8 alkyl, etc.; R3 = C1-8 alkyl, etc.; R4 = H, C1-4 alkyl; R5 = -R7-R8-R9- (R7 = C1-8 alkylene, R8 = methylene, imino, O, etc., and R9 = C1-10 alkylene, etc.); and R6 = H, C1-4 alkyl, provided that R1 and R3 in combination may form a ring. The compd. comprises a group I and any of **hyaluronic acid**, a deriv. thereof, and salts of these, the former being bonded to a hydroxyl group of the latter through a carbamate linkage'. Sodium **hyaluronate** was reacted with N-hydroxy-5-norbornene-2,3-dicarboxyimide (HONB) and hydroxamic acid deriv. N'-(13-amino-4,7,10-trioxatridecanyl)-N-(3S-hydroxy-4-(N-(1-methoxy-1-methylethoxy)amino)-2R-isobutylsuccinyl)-L-tert-leucinamide. The obtained compd. showed excellent inhibitory effect on gelatinase A and stromelysin-1 in in vitro test.

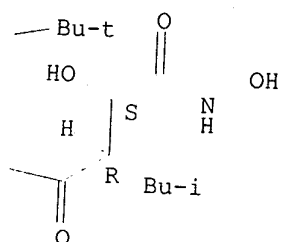
ST **hyaluronate** hydroxamate deriv prepn matrix metalloproteinase inhibitor

- IT Joint, anatomical  
(disease; **hyaluronic acid** hydroxamate derivs. for treatment of joint disease)
- IT Antiarthritics  
Antirheumatic agents  
(**hyaluronic acid** hydroxamate derivs. for treatment of joint disease)
- IT Collagens, biological studies  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(**hyaluronic acid** hydroxamate derivs. for treatment of joint disease)
- IT **434283-17-7DP**, complexes with **hyaluronic acid**  
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
(**hyaluronic acid** hydroxamate derivs. for treatment of joint disease)
- IT **434283-18-8D**, reaction products with **hyaluronate** derivs.  
**434283-19-9D**, reaction products with **hyaluronate** derivs.  
**434283-20-2D**, reaction products with **hyaluronate** derivs.  
**434283-21-3D**, reaction products with **hyaluronate** derivs.  
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(**hyaluronic acid** hydroxamate derivs. for treatment of joint disease)
- IT **79955-99-0**, Stromelysin-1 **141907-41-7**, Matrix metalloproteinase  
**146480-35-5**, Gelatinase A  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(inhibition of; **hyaluronic acid** hydroxamate derivs. for treatment of joint disease)
- IT **116-11-0** **5470-11-1**, Hydroxyammonium chloride **9067-32-7**, Sodium **hyaluronate** **21715-90-2**, HONb **62965-35-9**, N-(tert-Butoxycarbonyl)-L-tert-leucine **157518-70-2** **220156-99-0**  
RL: RCT (Reactant); RACT (Reactant or reagent)  
(prepn. of **hyaluronic acid** hydroxamate derivs. for treatment of joint disease)
- IT **433708-29-3P** **433708-31-7P** **433708-33-9P** **433708-35-1P**  
**433708-37-3P** **433708-39-5P**  
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
(prepn. of **hyaluronic acid** hydroxamate derivs. for treatment of joint disease)
- RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
- RE
- (1) Chugai Pharmaceutical Co Ltd; EP 1082963 A 1999 HCAPLUS  
(2) Chugai Pharmaceutical Co Ltd; WO 9959603 A 1999 HCAPLUS  
(3) Shionogi & Co Ltd; WO 0046189 A 2000 HCAPLUS
- IT **434283-17-7DP**, complexes with **hyaluronic acid**  
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
(**hyaluronic acid** hydroxamate derivs. for treatment of joint disease)
- RN **434283-17-7** HCAPLUS
- CN Carbamic acid, [3-[(18S,21R)-18-(1,1-dimethylethyl)-21-[(1S)-1-hydroxy-2-(hydroxyamino)-2-oxoethyl]-23-methyl-1,17,20-trioxo-6,9,12-trioxa-2,16,19-triazatetracos-1-yl]bicyclo[2.2.1]hept-5-en-2-yl]- (9CI) (CA INDEX NAME)
- Absolute stereochemistry.



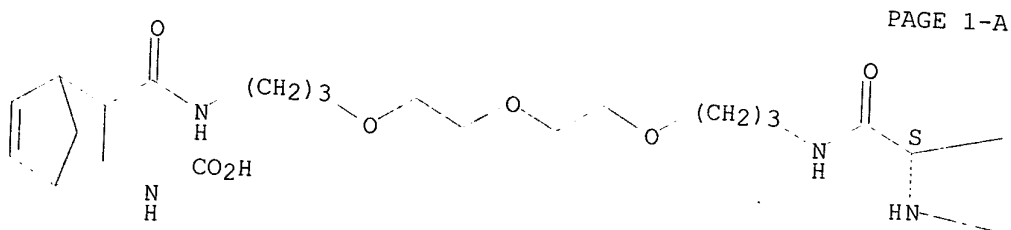


PAGE 1-B

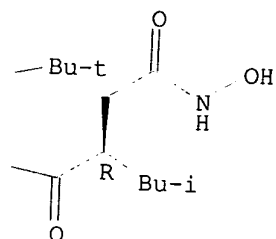


IT 434283-20-2D, reaction products with **hyaluronate** derivs.  
 434283-21-3D, reaction products with **hyaluronate** derivs.  
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
 (Biological study); USES (Uses)  
 (hyaluronic acid hydroxamate derivs. for treatment  
 of joint disease)  
 RN 434283-20-2 HCAPLUS  
 CN Carbamic acid, [3-[(18S,21R)-18-(1,1-dimethylethyl)-21-[2-(hydroxyamino)-2-  
 oxoethyl]-23-methyl-1,17,20-trioxo-6,9,12-trioxa-2,16,19-triazatetracos-1-  
 yl]bicyclo[2.2.1]hept-5-en-2-yl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



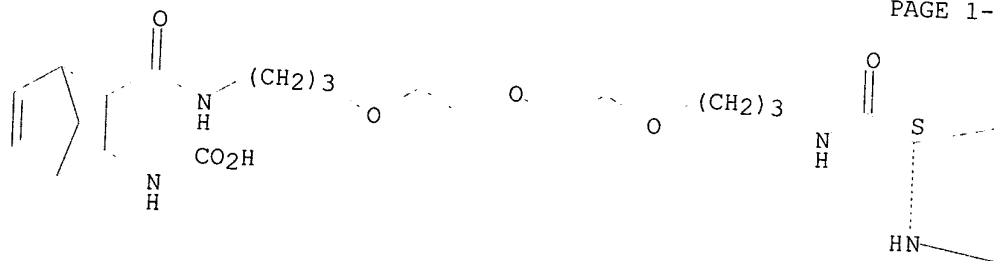
PAGE 1-B



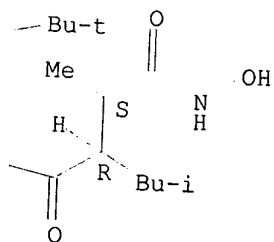
RN 434283-21-3 HCAPLUS  
 CN Carbamic acid, [3-[(18S,21R)-18-(1,1-dimethylethyl)-21-[(1S)-2-(hydroxyamino)-1-methyl-2-oxoethyl]-23-methyl-1,17,20-trioxo-6,9,12-trioxo-2,16,19-triazatetracos-1-yl]bicyclo[2.2.1]hept-5-en-2-yl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A



PAGE 1-B



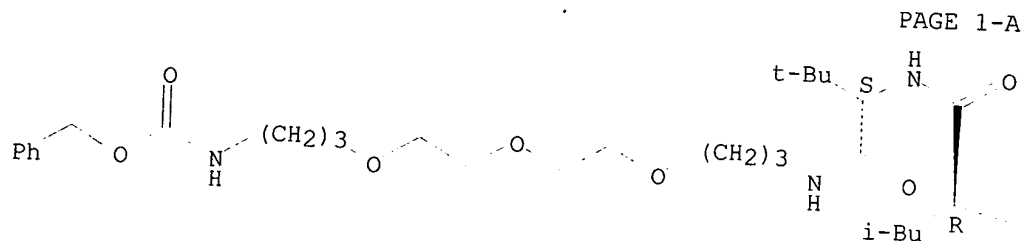
IT 9067-32-7, Sodium hyaluronate  
 RL: RCT (Reactant); RACT (Reactant or reagent)  
 (prepn. of hyaluronic acid hydroxamate derivs. for  
 treatment of joint disease)  
 RN 9067-32-7 HCAPLUS  
 CN Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

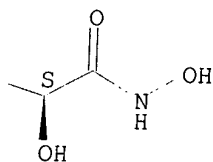
IT 433708-37-3p  
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT  
 (Reactant or reagent)  
 (prepn. of hyaluronic acid hydroxamate derivs. for  
 treatment of joint disease)

RN 433708-37-3 HCAPLUS  
 CN 6,9,12-Trioxa-2,16,19-triazatetracosanoic acid, 18-(1,1-dimethylethyl)-21-  
 [(1S)-1-hydroxy-2-(hydroxyamino)-2-oxoethyl]-23-methyl-17,20-dioxo-,  
 phenylmethyl ester, (18S,21R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



PAGE 1-B



L12 ANSWER 3 OF 4 HCAPLUS COPYRIGHT 2003 ACS  
 AN 2001:545502 HCAPLUS  
 DN 135:117219  
 TI Hapten-coagulation agent-antineoplastic agent combinations for treating  
 neoplasms  
 IN Yu, Baofa  
 PA USA  
 SO PCT Int. Appl., 83 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA English  
 IC ICM A61K033-40  
 ICS A61K031-06; A61K031-045; A61P035-00  
 CC 1-6 (Pharmacology)  
 Section cross-reference(s): 15  
 FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001052868	A1	20010726	WO 2001-US1737	20010118
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG US 2002044919 A1 20020418 PRAI US 2000-177024P P 20000119 US 2001-765060 20010117				

- AB Methods are provided for treating neoplasms, tumors and cancers, using one or more haptens and coagulation agents or treatments, alone or in combination with other anti-neoplastic agents or treatments. Also provided are combinations, and kits contg. the combinations for effecting the therapy.
- ST hapten coagulation agent antineoplastic agent combination antitumor
- IT Gene, animal  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(APC; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study) (B-lym; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(DCC; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study) (Ki-ras; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Cytokines  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(MBP (major basic protein); hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study) (N-myc; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study) (N-ras; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(NF-1; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(RB1; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(TP53; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(WT-1; hapten-coagulation agent-antineoplastic agent combinations for

- treating neoplasms)
- IT Adrenal cortex
  - (adrenocortical suppressants; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Interleukin 1
  - RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
  - (and anti-IL1 antibody; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Cytokines
  - RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
  - (and cytokine gene; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Chemokines
  - RL: BSU (Biological study, unclassified); BIOL (Biological study)
  - (angiostatic chemokine gene; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene
  - Steroids, biological studies
  - RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
  - (angiostatic; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Nutrients
  - (anti-; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antisense oligonucleotides
  - RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
  - (anti-oncogene; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Intestine, neoplasm
  - (anus, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents
  - (anus; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Nerve
  - (auditory, cancer inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Biliary tract
  - (bile duct, neoplasm, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents
  - (bladder carcinoma; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents
  - (bone; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents
  - (brain; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal
  - RL: BSU (Biological study, unclassified); BIOL (Biological study)
  - (c-Ha-ras; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal
  - RL: BSU (Biological study, unclassified); BIOL (Biological study)

- (c-abl; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal
  - RL: BSU (Biological study, unclassified); BIOL (Biological study)
  - (c-erbA; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal
  - RL: BSU (Biological study, unclassified); BIOL (Biological study)
  - (c-erbB; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal
  - RL: BSU (Biological study, unclassified); BIOL (Biological study)
  - (c-myc; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal
  - RL: BSU (Biological study, unclassified); BIOL (Biological study)
  - (c-sis; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Ear
  - Heart
  - Oviduct
  - Pituitary gland
  - Tonsil
  - (cancer inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Bladder
  - Esophagus
  - Kidney, neoplasm
  - Lung, neoplasm
  - Mammary gland
  - Ovary, neoplasm
  - (carcinoma, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Immunity
  - (cell-mediated; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents
  - (central nervous system; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Nervous system
  - (central, neoplasm, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Uterus, neoplasm
  - (cervix, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents
  - (cervix; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Intestine, neoplasm
  - (colon, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents
  - (colon; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Human immunodeficiency virus
  - (conditionally replicating, vector; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Therapy
  - (cryotherapy and transpupillary thermotherapy; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Cytolysis
  - (cytolytic gene; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

- IT Basement membrane  
(degrdn., inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(digestive tract; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Uterus, neoplasm  
(endometrium, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(endometrium; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Cytotoxic agents  
(endothelial cell; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Blood vessel  
(endothelium, endothelial cell proliferation inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(erbB2; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(esophagus carcinoma; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(esophagus; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(ets; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Brucella melitensis  
(ext.; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(eye; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(fes; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(fgr; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(fms; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(fos; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(fps; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Alkylating agents, biological  
Angiogenesis inhibitors  
Antitumor agents  
Chelating agents

Corynebacterium parvum  
 Coupling agents  
 Drug delivery systems  
 Immunostimulants  
 Immunotherapy  
 Mycobacterium BCG  
 Newcastle disease virus  
 Oxidizing agents  
 Radiosensitizers, biological  
 Radiotherapy  
 Reducing agents  
 Retroviral vectors  
 Surgery  
 Virus vectors  
 (hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)  
 IT Haptens  
 RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)  
 (hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)  
 IT Alcohols, biological studies  
 Antibodies  
 Enzymes, biological studies  
 Hormones, animal, biological studies  
 Interleukin 12  
 Interleukin 2  
 Interleukin 4  
 Laminins  
 Natural products  
 Ovalbumin  
 Polysaccharides, biological studies  
 Protamines  
 Reporter gene  
 Retinoids  
 Thrombospondins  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)  
 IT Antitumor agents  
 (head; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)  
 IT Liver, neoplasm  
 (hepatoma, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)  
 IT Antitumor agents  
 (hepatoma; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)  
 IT Herb  
 (herbal ext.; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)  
 IT Human herpesvirus  
 (herpes simplex viral amplicon vector; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)  
 IT Gene, animal  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)  
 (hit; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)  
 IT Gene, animal  
 RL: BSU (Biological study, unclassified); BIOL (Biological study)



- (hst; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Immunity
  - (humoral; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Adrenal gland, neoplasm
- Bone, neoplasm
- Brain, neoplasm
- Cell migration
- Eye, neoplasm
- Kidney, neoplasm
- Lung, neoplasm
- Ovary, neoplasm
- Pancreas, neoplasm
- Skin, neoplasm
- Stomach, neoplasm
- Testis, neoplasm
- Thyroid gland, neoplasm
- Uterus, neoplasm
  - (inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Drug delivery systems
  - (injections; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal
  - RL: BSU (Biological study, unclassified); BIOL (Biological study) (int-1; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal
  - RL: BSU (Biological study, unclassified); BIOL (Biological study) (int2; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Proteins, specific or class
  - RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (interferon .gamma.-inducible protein 10; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal
  - RL: BSU (Biological study, unclassified); BIOL (Biological study) (jun; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents
  - (kidney carcinoma; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents
  - (kidney; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents
  - (larynx tumor inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Lasers
  - (laser coagulation; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Eye
  - (lid, cancer inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents
  - (lung carcinoma; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents
  - (lung non-small-cell carcinoma; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Antitumor agents  
(lung; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Antitumor agents  
(mammary gland carcinoma; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Antitumor agents  
(mammary gland; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Jaw  
(mandibula, cancer inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Jaw  
(mandibula, condylar process, cancer inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(mas; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Jaw  
(maxilla, cancer inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(met; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(mil; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(mos; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Antitumor agents  
(mouth; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(myb; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Pharynx  
(nasopharynx, neoplasm, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Antitumor agents  
(nasopharynx; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Antitumor agents  
(neck; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT Digestive tract  
Esophagus  
Head  
Mammary gland  
Mouth  
Neck, anatomical  
Nose  
Prostate gland  
Salivary gland  
Spinal cord  
Urethra  
(neoplasm, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(neu; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT *Vibrio cholerae*  
(neuraminidase; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Lung, neoplasm  
(non-small-cell carcinoma, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Virus  
(nonvirulent; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(oncogene, inhibitor; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(ovary carcinoma; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(ovary; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(p16; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(p21; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(p27; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(pancreas; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Salivary gland  
(parotid, cancer inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(penis tumor inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Fibronectins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(peptides; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Microwave  
(percutaneous microwave coagulation therapy; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Proteins, specific or class  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(placental proliferin-related protein; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

- IT Proliferation inhibition  
(proliferation inhibitors, endothelial cell; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Proteins, specific or class  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(proliferin-related protein; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(prostate gland; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Denaturants  
(protein denaturing agents; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Denaturation  
(protein, agents for; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Necrosis  
(radio-frequency-induced coagulation necrosis; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(raf; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(ral; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Intestine, neoplasm  
(rectum, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(rectum; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(rel; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Eye  
(retina, cancer inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(ros; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(ski; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(skin; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(small intestine; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Intestine, neoplasm  
(small, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(solid tumor; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents

- (spinal cord; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(src; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(stomach; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(suicide gene; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(testis; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(thyroid; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(trk; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Larynx  
Penis  
(tumor inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Proteins, specific or class  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(tumor suppressor protein; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(tumor suppressor; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Vagina  
(tumor, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antigens  
RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
(tumor-assocd.; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Fibroblast growth factor receptors  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(type 1, sol.; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Sound and Ultrasound  
(ultrasonic therapy; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(urethra; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(uterus; hapten-coagulation agent-antineoplastic agent combinations for

- treating neoplasms)
- IT Immunization  
(vaccination; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(vaginal tumor inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Adenoviridae  
Simian virus 40  
Vaccinia virus  
(vector; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Nerve  
(vestibulocochlear, cancer inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Fluids  
(vitreous; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Reproductive tract  
(vulva, neoplasm, inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Antitumor agents  
(vulva; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Gene, animal  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(yes; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Tumor necrosis factors  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(.alpha., antibody to; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Interferons  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(.alpha.; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Integrins  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(.alpha.v.beta.3, antibody to; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT Interferons  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(.gamma.; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT 9001-67-6, Neuraminidase  
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(Vibrio cholera; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT 127464-60-2, Vascular endothelial growth factor  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(antibody to, and VEGF inhibitors; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT 106096-93-9, Basic fibroblast growth factor  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(antibody to; hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)
- IT 50-01-1, Guanidine hydrochloride 50-02-2, Dexamethasone 50-18-0,

Cyclophosphamide 50-23-7, Hydrocortisone 50-24-8, Prednisolone 52-67-5, D-Penicillamine 53-02-1, Tetrahydrocortisol 53-06-5, Cortisone 53-86-1, Indomethacin 54-05-7, Chloroquine 56-81-5, Glycerol, biological studies 57-13-6, Urea, biological studies 57-13-6D, Urea, derivs., biological studies 57-55-6, 1,2-Propanediol, biological studies 58-27-5, Menadione 59-05-2, Methotrexate 60-24-2, 2-Mercaptoethanol 60-34-4D, Methylhydrazine, derivs. 64-17-5, Ethyl alcohol, biological studies 67-56-1, Methyl alcohol, biological studies 67-63-0, Isopropyl alcohol, biological studies 67-66-3, Chloroform, biological studies 70-34-8, Dinitrofluorobenzene 71-23-8, n-Propyl alcohol, biological studies 71-36-3, n-Butyl alcohol, biological studies 71-41-0, n-Pentyl alcohol, biological studies 75-65-0, tert-Butyl alcohol, biological studies 75-85-4, tert-Pentyl alcohol 75-91-2, tert-Butyl hydroperoxide 78-83-1, Isobutyl alcohol, biological studies 78-92-2, sec-Butyl alcohol 88-89-1, Trinitrophenol 96-41-3, Cyclopentanol 104-54-1, Cinnamyl alcohol 107-18-6, Allyl alcohol, biological studies 107-21-1, 1,2-Ethanediol, biological studies 108-93-0, Cyclohexanol, biological studies 108-95-2, Phenol, biological studies 111-27-3, n-Hexyl alcohol, biological studies 111-70-6, n-Heptyl alcohol 111-87-5, n-Octyl alcohol, biological studies 112-30-1, n-Decyl alcohol 112-53-8, n-Dodecyl alcohol 112-72-1, n-Tetradecyl alcohol 112-92-5, n-Octadecyl alcohol 115-77-5, Pentaerythritol, biological studies 123-51-3, Isopentyl alcohol 128-08-5, N-Bromosuccinimide 128-53-0, N-Ethylmaleimide 137-32-6, Active-amyl alcohol 145-63-1, Suramin 147-94-4, AraC 151-51-9, Carbodiimide 152-58-9, Cortexolone 342-69-8, 6-Methylmercaptopurine riboside 446-86-6, Azathioprine 504-63-2, 1,3-Propanediol 517-28-2, Hematoxylin 520-85-4, Medroxyprogesterone 593-84-0, Guanidinium thiocyanate 994-36-5, Sodium citrate 1398-61-4, Chitin 4846-27-9 6117-91-5, Crotyl alcohol 7440-06-4D, Platinum, coordination complexes, biological studies 7585-39-9, .beta.-Cyclodextrin 7722-84-1, Hydrogen peroxide, biological studies 7790-28-5, Sodium periodate 8049-47-6, Pancreatin 9001-73-4, Papain 9002-62-4D, Prolactin, 16-kDa fragment, biological studies 9004-61-9, Hyaluronan 9005-49-6, Heparin, biological studies 9012-72-0, Glucan 9025-39-2, Heparinase 10028-15-6, Ozone, biological studies 10102-43-9, Nitric oxide, biological studies 10118-90-8, Minocycline 10361-76-9, Potassium peroxymonosulfate 10465-78-8, Diamide 11103-57-4, vitamin A 11118-27-7, Gold chloride 14769-73-4, Levamisole 15307-86-5, Diclofenac 15663-27-1, Cisplatin 15687-27-1, Ibuprofen 15866-90-7, Metastat 22668-01-5, SR 2508 23214-92-8D, Doxorubicin, conjugates with adipic dihydrazide 25550-58-7, Dinitrophenol 27314-97-2, Tirapazamine 27591-97-5, Tilorone 33069-62-4, Paclitaxel 33507-63-0, Substance P 34031-32-8, Auranoftin 36653-82-4, 1-Hexadecanol 36877-68-6D, Nitroimidazole, derivs. 36930-63-9 37270-94-3, platelet factor 4 39450-01-6 51110-01-1, Somatostatin 51592-06-4, Iodogen 59865-13-3, Cyclosporin A 73590-58-6, Omeprazole 75706-12-6, SU101 83150-76-9, Octreotide 83869-56-1, GM-CSF 84088-42-6, Linomide 86090-08-6, Angiostatin 105844-41-5, Plasminogen activator inhibitor 108121-76-2D, Anthracenedione, derivs. 124861-55-8 126857-36-1, 08, biological studies 129298-91-5, AGM-1470 130370-60-4, BB-94 134633-29-7, Tecogalan sodium 140207-93-8 140208-24-8, tissue inhibitor of metalloproteinase-1 145809-21-8, tissue inhibitor of metalloproteinase-3 148805-91-8 153851-75-3, Heptoxepane 154039-60-8, BB-2516 166981-13-1, CT-2584 184110-80-3, GM 1474 188417-67-6, CM 101 203515-84-8 324740-00-3, Vitaxin

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(hapten-coagulation agent-antineoplastic agent combinations for treating neoplasms)

IT 9040-48-6, Gelatinase 9055-65-6, prostaglandin synthase 79955-99-0, Stromelysin 1 141907-41-7, Matrix metalloproteinase

RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(inhibitor; hapten-coagulation agent-antineoplastic agent combinations  
for treating neoplasms)

IT 9001-99-4, RNase

RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(placental RNase inhibitor; hapten-coagulation agent-antineoplastic  
agent combinations for treating neoplasms)

RE.CNT 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD

RE

(1) Battentier, E; FR 2505182 A 1982 HCAPLUS

(2) Berd, D; US 5290551 A 1994 HCAPLUS

(3) Cone, C; US 4724230 A 1988 HCAPLUS

(4) du Pont; EP 0378888 A 1990 HCAPLUS

(5) Roy, W; WO 0006143 A 2000 HCAPLUS

(6) Rubin, D; US 5005588 A 1991

(7) Rupchock, P; US 4447526 A 1984 HCAPLUS

(8) Zhang, M; Melanoma Research 1998, V8(6), P510 HCAPLUS

IT 9004-61-9, Hyaluronan 154039-60-8, BB-2516

RL: BAC (Biological activity or effector, except adverse); BSU (Biological  
study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES  
(Uses)

(hapten-coagulation agent-antineoplastic agent combinations for  
treating neoplasms)

RN 9004-61-9 HCAPLUS

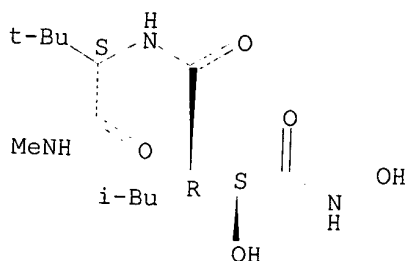
CN Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

RN 154039-60-8 HCAPLUS

CN Butanediamide, N4-[(1S)-2,2-dimethyl-1-[(methylamino)carbonyl]propyl]-N1,2-  
dihydroxy-3-(2-methylpropyl)-, (2S,3R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 4 OF 4 HCAPLUS COPYRIGHT 2003 ACS

AN 1994:46 HCAPLUS

DN 120:46

TI A stromelysin assay for the assessment of metalloprotease inhibitors on  
human aggregated proteoglycan

AU Doughty, J. R.; Goldberg, R. L.; Ganu, V.; Melton, R. A.; Hu, S. I.; Di  
Pasquale, G.

CS Pharm. Div., CIBA-GEIGY Corp., Summit, NJ, 07901, USA

SO Agents and Actions (1993), 39(Spec. Conf. Issue), C151-C153

CODEN: AGACBH; ISSN: 0065-4299

DT Journal

LA English

CC 1-1 (Pharmacology)

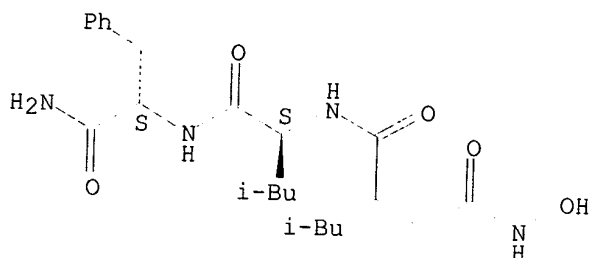
AB Human proteoglycan was aggregated to an immobilized **hyaluronan**  
solid phase on a 96-well ELISA plate. This complex was then degraded by  
recombinant human stromelysin. The remaining proteoglycan fragments were  
detected using a monoclonal antibody probe directed against the  
chondroitin sulfate (CS) region of the core protein. Stromelysin degraded



the aggregate in a time and dose dependent manner as reflected by the loss of the CS epitope. Assay sensitivity was 0.125 U/well with total loss of the CS epitope occurring at 4 U/well. O-phenanthroline (IC50 = 52 .mu.M) and U24522 (IC50 = 9 .mu.M) inhibited degrdn., while phosphoramidon did not. Serine and cysteine protease inhibitors had no effect. A comparative anal. of this assay with a ref. method, substance P assay, gave similar inhibitor profiles. The use of aggregated human proteoglycan (native conformation) as a substrate, may better reflect how stromelysin inhibitors behave in the presence of complex substrates such as cartilage matrix.

- ST stromelysin assay metalloprotease inhibitor aggregated proteoglycan  
 IT Inflammation inhibitors  
     (antiarthritics, metalloprotease inhibitors as, proteoglycan degrdn. inhibition as assay of)  
 IT Proteoglycans, biological studies  
     RL: PRP (Properties)  
     (chondroitin sulfate-contg., metalloprotease inhibitors prevention of degrdn. of, by stromelysin, antiarthritics assay by)  
 IT 79955-99-0, Stromelysin  
     RL: BSU (Biological study, unclassified); BIOL (Biological study)  
     (inhibitors, assay of antiarthritic activity of, proteoglycan degrdn. inhibition in)  
 IT 66-71-7, o-Phenanthroline **106314-87-8**, U24522  
     RL: ANST (Analytical study)  
     (proteoglycan degrdn. inhibition by, as metalloprotease inhibitor, antiarthritic activity in relation to)  
 IT 36357-77-4, Phosphoramidon  
     RL: ANST (Analytical study)  
     (proteoglycan degrdn. response to, as metalloprotease inhibitor, antiarthritic activity in relation to)  
 IT **106314-87-8**, U24522  
     RL: ANST (Analytical study)  
     (proteoglycan degrdn. inhibition by, as metalloprotease inhibitor, antiarthritic activity in relation to)  
 RN 106314-87-8 HCAPLUS  
 CN L-Phenylalaninamide, N-[2-[2-(hydroxyamino)-2-oxoethyl]-4-methyl-1-oxopentyl]-L-leucyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



=> fil reg

FILE 'REGISTRY' ENTERED AT 16:41:48 ON 27 JAN 2003  
 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.  
 PLEASE SEE "HELP USAGETERMS" FOR DETAILS.  
 COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 26 JAN 2003 HIGHEST RN 481631-75-8

DICTIONARY FILE UPDATES: 26 JAN 2003 HIGHEST RN 481631-75-8

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

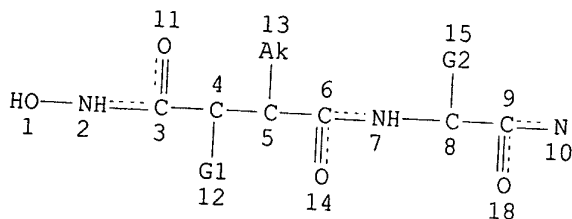
Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details:  
<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

=> d sta que 15

L3

STR

Ak~Cy  
 @16 17



VAR G1=H/OH/AK

VAR G2=AK/16

NODE ATTRIBUTES:

CONNECT IS M1 RC AT 10

DEFAULT MLEVEL IS ATOM

DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED

NUMBER OF NODES IS 18

STEREO ATTRIBUTES: NONE

L4 ( 2264)SEA FILE=REGISTRY SSS FUL L3

L5 629 SEA FILE=REGISTRY SUB=L4 CSS FUL L3

100.0% PROCESSED 2264 ITERATIONS

SEARCH TIME: 00.00.01

629 ANSWERS